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Recipient: IB

- 30. The method of Claim 29, wherein the first cistron encodes CREB variant Y134F.
- 31. The method of Claim 22, wherein the first cistron encodes a CREB protein or a variant thereof, and the second cistron encodes a Bcl-2 protein or a Bcl-2 protein having a deletion in the regulatory loop domain.
- The method of Claim 22, wherein the first cistron encodes a variant E1a protein with a mutation in CR1, and the second cistron encodes an E1b-19K protein, a Bcl-2 protein, or a Bcl-2 protein having a deletion in the regulatory loop domain.
  - 33. The method of Claim 22, wherein the second cistron encodes an apoptosis-protective protein selected from the group consisting of a dominant negative mutant of p53, a protein that interacts with BAX, a protein that interacts with BAK, an inhibitor of apoptosome formation, and a downstream apoptosis inhibitor.

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- 34. The method of Claim 22, wherein the second cistron encodes an adenovirus E1b-19K protein, a Bcl-2 protein, or a Bcl-2 protein having a deletion in the regulatory loop domain.
- The method of Claim 22, wherein said polypeptide is a single-chain antibody or a heavy or light chain of an antibody or antibody fragment.
  - 36. The method of Claim 22, wherein said polypeptide is a part of a library of polypeptides.
  - 37. A mammalian host cell for recombinant polypeptide expression comprising a first cistron encoding a transactivator protein and a second cistron encoding an apoptosis-protective protein that prevents cell-killing due to expression of the transactivator protein.
    - 38. The host cell of Claim 37, further comprising a third cistron encoding one or more desired polypeptide under the control of a promoter responsive to the transactivator protein.
    - 39. The host cell of Claim 37, wherein the third cistron is associated with a ubiquitous chromatin opening element, an insulator, or a barrier element.
    - 40. The host cell of Claim 37, wherein the transactivator protein is expressed from an efficient heterologous promoter.

MAQAGRTGYDNREIVMKYIHYKLSQRGYEWDVGDVDAAPLG
AAPTPGIFSFQPESNPTPAVHRDMAARTSPLRPIVATTGPT
LSPVPPVVHLTLRRAGDDFSRRYRRDFAEMSSQLHLTPFTA
RGRFATVVEELFRDGVNWGRIVAFFEFGGVMCVESVNREMS
PLVDNIALWMTEYLNRHLHTWIQDNGGWDAFVELYGPSVRP
LFDFSWLSLKTLLSLALVGACITLGTYLGHK (SEQ
ID NO: 1)

FIG. 1

MAQAGRTGYDNREIVMKYIHYKLSQRGYEWDVGDVDAAAAA
ASPVPPVVHLTLRRAGDDFSRRYRRDFAEMSSQLHLTPFTA
RGRFATVVEELFRDGVNWGRIVAFFEFGGVMCVESVNREMS
PLVDNIALWMTEYLNRHLHTWIQDNGGWDAFVELYGPSVRP
LFDFSWLSLKTLLSLALVGACITLGTYLGHK (SEQ
ID NO: 2)

FIG. 2

1 ATGGCTCAAG CTGGGAGAAC AGGGTATGAT AACCGAGAGA TCGTGATGAA
51 GTACATCCAT TATAAGCTGT CACAGAGGGG CTACGAGTGG GATGTGGGAG
101 ATGTGGACGC CGCGGCCGCG GCCGCGAGCC CCGTGCCACC TGTGGTCCAC
151 CTGACCCTCC GCCGGGCTGG GGATGACTTC TCCCGTCGCT ACCGTCGCGA
201 CTTCGCGGAG ATGTCCAGTC AGCTGCACCT GACGCCCTTC ACCGCGAGGG
251 GACGCTTTGC TACGGTGGTG GAGGAACTCT TCAGGGATGG GGTGAACTGG
301 GGGAGGATTG TGGCCTTCTT TGAGTTCGGT GGGGTCATGT GTGTGGAGAG
351 CGTCAACAGG GAGATGTCAC CCCTGGTGGA CAACATCGCC CTGTGGATGA
401 CCGAGTACCT GAACCGGCAT CTGCACACCT GGATCCAGGA TAACGGAGGC
451 TGGGACGCAT TTGTGGAACT GTACGGCCCC AGTGTGAGGC CTCTGTTTGA
501 TTTCTCTTGG CTGTCTCTGA AGACCCTGCT CAGCCTGGCC CTGGTCGGGG
551 CCTGCATCAC TCTGGGTACC TACCTGGGCC ACAAGTGA (SEQ ID NO: 3)

FIG. 3

MRHIICHGGVITEEMAASLLDQLIEEVLADNLPPPSHFE
PPTLHELHDLDVTAPEDPNEEAVSQIFPDSVMLAVQEGI
DLLTFPPAPGSPEPPHLSRQPEQPEQRALGPVSMPNLVP
EVIDLTGHEAGFPPSDDEDEEGEEFVLDYVEHPGHGCR
SCHYHRRNTGDPDIMCSLCYMRTCGMFVYSPVSEPEPE
PEPEPARPTRRPKMAPAILRRPTSPVSRECNSSTDSCD
SGPSNTPPEIHPVVPLCPIKPVAVRVGGRRQAVECIEDL
LNEPGQPLDLSCKRPRP (SEQ ID NO: 4)

FIG. 4

1 ATGAGACATA TTATCTGCCA CGGAGGTGTT ATTACCGAAG AAATGGCCGC 51 CAGTCTTTTG GACCAGCTGA TCGAAGAGGT ACTGGCTGAT AATCTTCCAC 101 CTCCTAGCCA TTTTGAACCA CCTACCCTTC ACGAACTGCA TGATTTAGAC 151 GTGACGGCCC CCGAAGATCC CAACGAGGAG GCGGTTTCGC AGATTTTTCC 201 CGACTCTGTA ATGTTGGCGG TGCAGGAAGG GATTGACTTA CTCACTTTTC 251 CGCCGCCCC CGGTTCTCCG GAGCCGCCTC ACCTTTCCCG GCAGCCCGAG 301 CAGCCGGAGC AGAGAGCCTT GGGTCCGGTT TCTATGCCAA ACCTTGTACC 351 GGAGGTGATC GATCTTACCG GCCACGAGGC TGGCTTTCCA CCCAGTGACG 401 ACGAGGATGA AGAGGGTGAG GAGTTTGTGT TAGATTATGT GGAGCACCCC 451 GGGCACGGTT GCAGGTCTTG TCATTATCAC CGGAGGAATA CGGGGGACCC 501 AGATATTATG TGTTCGCTTT GCTATATGAG GACCTGTGGC ATGTTTGTCT 551 ACAGTAAGTG AAAATTATGG GCAGTGGGTG ATAGAGTGGT GGGTTTGGTG 601 TGGTAATTTT TTTTTAATT TTTACAGTTT TGTGGTTTAA AGAATTTTGT 651 ATTGTGATTT TTTTAAAAGG TCCTGTGTCT GAACCTGAGC CTGAGCCCGA 701 GCCAGAACCG GAGCCTGCAA GACCTACCCG CCGTCCTAAA ATGGCGCCTG 751 CTATCCTGAG ACGCCCGACA TCACCTGTGT CTAGAGAATG CAATAGTAGT 801 ACGGATAGCT GTGACTCCGG TCCTTCTAAC ACACCTCCTG AGATACACCC 851 GGTGGTCCCG CTGTGCCCCA TTAAACCAGT TGCCGTGAGA GTTGGTGGGC 901 GTCGCCAGGC TGTGGAATGT ATCGAGGACT TGCTTAACGA GCCTGGGCAA 951 CCTTTGGACT TGAGCTGTAA ACGCCCCAGG CCATAA (SEQ ID NO: 5)

FIG. 5

1 GAATTCGCCG CCACCATGGA GGCTTGGGAG TGTTTGGAAG ATTTTCTGC
51 TGTGCGTAAC TTGCTGGAAC AGAGCTCTAA CAGTACCTCT TGGTTTTGGA
101 GGTTTCTGTG GGGCTCATCC CAGGCAAAGT TAGTCTGCAG AATTAAGGAG
151 GATTACAAGT GGGAATTTGA AGAGCTTTTG AAATCCTGTG GTGAGCTGTT
201 TGATTCTTTG AATCTGGGTC ACCAGGCGCT TTTCCAAGAG AAGGTCATCA
251 AGACTTTGGA TTTTTCCACA CCGGGGCGCG CTGCGGCTGC TGTTGCTTTT
301 TTGAGTTTTA TAAAGGATAA ATGGAGCGAA GAAACCCATC TGAGCGGGGG
351 GTACCTGCTG GATTTTCTGG CCATGCATCT GTGGAGAGCG GTTGTGAGAC
401 ACAAGAATCG CCTGCTACTG TTGTCTTCCG TCCGCCCGGC GATAATACCG
451 ACGGAGGAGC AGCAGCAGCA GCAGGAGGAA GCCAGGCGGC GGCGGCAGGA
501 GCAGAGCCCA TGGAACCCGA GAGCCGGCCT GGACCCTCGG GAATGAATGT
551 TGGTCGAC (SEQ ID NO: 15)

FIG. 6

- 1 GTCGACGCCG CCACCATGCC GCCCAAAACC CCCCGAAAAA CGGCCGCCAC
  M P P K T P R K T A A T
- 51 CGCCGCCGCT GCCGCCGCGG AACCCCCGGC ACCGCCGCCG CCGCCCCCTC
  A A A A A A A B P P A P P P P P P P
- 101 CTGAGGAGGA CCCAGAGCAG GACAGCGGCC CGGAGGACCT GCCTCTCGTC
  E E D P E Q D S G P E D L P L V
- 151 AGGCTTGAGT TTGAAGAAAC AGAAGAACCT GATTTTACTG CATTATGTCA R L E F E E T E E P D F T A L C O
- 201 GAAATTAAAG ATACCAGATC ATGTCAGAGA GAGAGCTTGG TTAACTTGGG K L K I P D H V R E R A W L T W E
- 251 AGAAAGTTTC ATCTGTGGAT GGAGTATTGG GAGGTTATAT TCAAAAGAAA K V S S V D G V L G G Y I Q K K
- 301 AAGGAACTGT GGGGAATCTG TATCTTTATT GCACGAGTTG ACCTAGATGA
  K E L W G I C I F I A R V D L D E
- 351 GATGTCGTTC ACTTTACTGA GCTACAGAAA AACATACGAA ATCAGTGTCC M S F T L L S Y R K T Y E I S V H
- 401 ATAAATTCTT TAACTTACTA AAAGAAATTG ATACCAGTAC CAAAGTTGAT K F F N L L K E I D T S T K V D
- 451 AATGCTATGT CAAGACTGTT GAAGAAGTAT GATGTATTGT TTGCACTCTT N A M S R L L K K Y D V L F A L F
- 501 CAGCAAATTG GAAAGGACAT GTGAACTTAT ATATTTGACA CAACCCAGCA S K L E R T C E L I Y L T Q P S S
- 551 GTTCGATATC TACTGAAATA AATTCTGCAT TGGTGCTAAA AGTTTCTTGG S I S T E I N S A L V L K V S W
- 601 ATCACATTT TATTAGCTAA AGGGGAAGTA TTACAAATGG AAGATGATCT I T F L L A K G E V L Q M E D D L
- 651 GGTGATTTCA TTTCAGTTAA TGCTATGTGT CCTTGACTAT TTTATTAAAC
  V I S F Q L M L C V L D Y F I K L

FIG. 7A

701 TCTCACCTCC CATGTTGCTC AAAGAACCAT ATAAAACAGC TGTTATACCC S P P M L L K E P Y K T A V I P 751 ATTAATGGTT CACCTCGAAC ACCCAGGCGA GGTCAGAACA GGAGTGCACG INGS PRT PRR G Q N R 801 GATAGCAAAA CAACTAGAAA ATGATACAAG AATTATTGAA GTTCTCTGTA IAK Q L E N D T R I I E V L C K 851 AAGAACATGA ATGTAATATA GATGAGGTGA AAAATGTTTA TTTCAAAAAT E H E C N I D E V K N V Y F K N ECORI 901 TTTATACCTT TTATGAATTC TCTTGGACTT GTAACATCTA ATGGACTTCC F I P F M N S L G L V T S N G L P 951 AGAGGTTGAA AATCTTTCTA AACGATACGA AGAAATTTAT CTTAAAAATA EVENLSKRYEEIYLKNK 1001 AAGATCTAGA TCGAAGATTA TTTTTGGATC ATGATAAAAC TCTTCAGACT D L D R R L F L D H D K T L O T 1051 GATTCTATAG ACAGTTTTGA AACACAGAGA ACACCACGAA AAAGTAACCT D S I D S F E T Q R T P R K S N L 1101 TGATGAAGAG GTGAATATAA TTCCTCCACA CACTCCAGTT AGGACTGTTA DEE VNII PPH TPV RTV M 1151 TGAACACTAT CCAACAATTA ATGATGATTT TAAATTCTGC AAGTGATCAA N T I Q Q L M M I L N S A S D O 1201 CCTTCAGAAA ATCTGATTTC CTATTTTAAC AACTGCACAG TGAATCCAAA PSENLIS Y FN N C T V N P K 1251 AGAAAGTATA CTGAAAAGAG TGAAGGATAT AGGATACATC TTTAAAGAGA ESI L K R V K D I G Y I F K E K 1301 AATTTGCTAA AGCTGTGGGA CAGGGTTGTG TCGAAATTGG ATCACAGCGA F A K A V G O G C V E I G S O R 1351 TACAAACTTG GAGTTCGCTT GTATTACCGA GTAATGGAAT CCATGCTTAA Y K L G V R L Y Y R V M E S M L K

FIG. 7B

1401				TTTTAGCAAA	
	S E E	E R L S	I Q N	F S K	r r n d
1451				GCGCTCTTGA A L E	
1501	GCCACATATA A T Y S			CTTGATTCTG L D S G	GAACAGATTT T D L
1551				TTTAAAAGCC L K A	
1601				AAGGCAACTT G N L	
1651		ATTTAGAACG L E R		CGAATCATGG R I M E	AATCCCTTGC S L A
1701				TATTAAACAA I K Q	
1751	GAGAAGGACC E G P			CTTGTCCTCT C P L	
1801				TATCTTTCTC Y L S P	
1851		AAAGGTTCAA K G S T		AAATTCTACT N S T	GCAAATGCAG A N A E
1901				AGAAGCCATT K P L	
1951				CGGCTAGCCT R L A Y	
2001				GCACCCAGAA H P E	
2051		CCTTTTCCAG L F Q		AGAATGAGTA N E Y	

FIG. 7C

2101	AGAGACAGGC				
	RDRH	L D Q	I M M	C S M Y	GIC
2151				AATCATTGTA I I V	
2201				TCAAACGTGT K R V	
2251				TATAACTCGG Y N S V	
2301	GAGACTGAAA R L K			TTCCACCAGG S T R	
2351				CTTACAAGTT Y K F	
2401				ATTTCACCCC I S P L	
2451				AACAAAAATG T K M	
2501				TCGGGACTTC G T S	
2551				GACCGTGTGC D R V L	
2601				GAAAAAACTA K K L	
2651				AACATCTCCC H L P	
2701				TCTACTCGAA S T R T	
2751				CTCAAACAAG S N K	

NotI 2801 GAGGATCTCA GGACCGGCGG CCGC

FIG. 7D

EcoRI 1 GAATTCGCCG CCACCATGAC CATGGACTCT GGAGCAGACA ACCAGCAGAG M T M D S G A D N Q Q S 51 TGGAGATGCA GCTGTAACAG AAGCTGAAAA CCAACAAATG ACAGTTCAAG G D A A V T E A E N Q Q M T V O A 101 CCCAGCCACA GATTGCCACA TTAGCCCAGG TATCTATGCC AGCAGCTCAT O P O I A T L A O V S M P A A H 151 GCAACATCAT CTGCTCCCAC CGTAACTCTA GTACAGCTGC CCAATGGGCA ATSSAPT V T L V O L P N G O 201 GACAGTTCAA GTCCATGGAG TCATTCAGGC GGCCCAGCCA TCAGTTATTC T V O V H G V I O A A O P S V I O 251 AGTCTCCACA AGTCCAAACA GTTCAGATTT CAACTATTGC AGAAAGTGAA S P Q V Q T V Q I S T I A E S E 301 GATTCACAGG AGTCAGTGGA TAGTGTAACT GATTCCCAAA AGCGAAGGGA D S Q E S V D S V T D S Q K R R E 351 AATTCTTTCA AGGAGGCCTT CCTTCAGGAA AATTTTGAAT GACTTATCTT ILS RRPS FRKILN DLSS 401 CTGATGCACC AGGAGTGCCA AGGATTGAAG AAGAGAAGTC TGAAGAGGAG DAPGVPRIEE EKSEEE 451 GCTTCAGCAC CTGCCATCAC CGCTGTAGCG GTGCCAACGC CAATTTACCG A S A P A I T A V A V P T P I Y R 501 GACTAGCAGT GGACAGTATA TTACCATTAC CCAGAGAGGA GCAATACAGC T S S G O Y I T I T O R G A I O L 551 TGGCTAGCAA TGGTACCGAT GGGGTACAGG GCCTGCAAAC ATTAACCATG A S N G T D G V O G L O T L T M 601 GCCAATGCAG CAGCCACTCA GCCGGGTACT ACCATTCTAC AGTATGCACA ANAA ATQ PGT TILQ YAQ 651 GACCACTGAT GGACAGCAGA TCTTAGTGCC CAGCAACCAA GTTGTTGTTC T T D G Q Q I L V P S N O V V V Q

## FIG. 8A

- 701 AAGCTGCCTC TGGAGACGTA CAAACATACC AGATTCGCAC AGCACCCACT A A S G D V Q T Y Q I R T A P T

  751 AGCACTATTG CCCCTGGAGT TGTTATGGCA TCCTCCCCAG CACTTCCTAC S T I A P G V V M A S S P A L P T

  801 ACAGCCTGCT GAAGAAGCAG CACGAAAGAG AGAGGTCCGT CTAATGAAGA Q P A E E A A R K R E V R L M K N

  851 ACAGGGAAGC AGCTCGTGAG TGTCGTAGAA AGAAGAAAGA ATATGTGAAA R E A A R E C R R K K K E Y V K
- 901 TGTTTAGAAA ACAGAGTGGC AGTGCTTGAA AATCAAAACA AGACATTGAT C L E N R V A V L E N Q N K T L I
- 951 TGAGGAGCTA AAAGCACTTA AGGACCTTTA CTGCCACAAA TCAGATTAAT E E L K A L K D L Y C H K S D \*

SalI 1001 TTGGGTCGAC

FIG. 8B

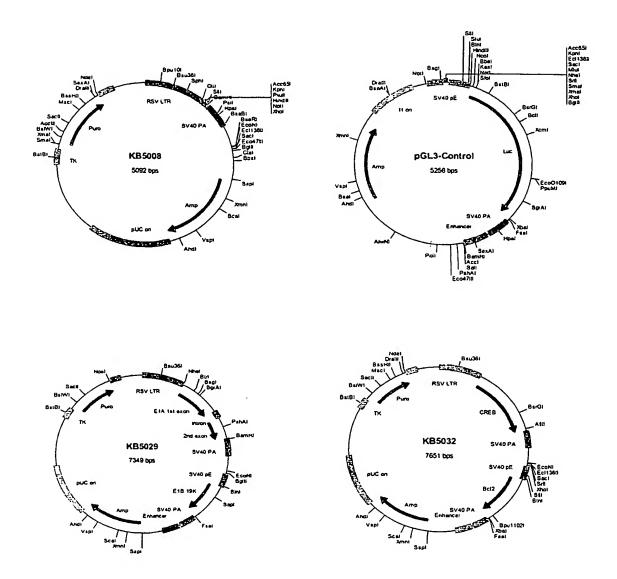


Figure 9. Maps of KB5008, pGL3-Control, KB5029, and KB5032.

KB5008 carries a RSVLTR promoter and was used to clone E1a and CREB. pGL3-Control carries a SV40 promoter and was used to clone E1b-19K and Bcl2. KB5029 carries the double expression cassettes for E1a and E1b-19K. KB5032 carries the double expression cassettes for CREB and Bcl2.

```
aagettgeeg eeaceatgag acatattate tgeeacggag gtgttattae HIROLIII M R H I I C H G G V I
       cgaagaaatg gccgccagtc ttttggacca gctgatcgaa gaggtactgg T E E M A A S L L D Q L I E E V L
        ctgataatct tccacctcct agccattttg aaccacctac ccttcacgaa A D N L P P P S H F E P P T L H E
101
       ctgtatgatt tagacgtgac ggccccgaa gatcccaacg aggaggcggt L \underline{Y} D L D \underline{V} T \underline{A} P \underline{E} D \underline{P} \underline{N} \underline{E} \underline{E} \underline{A}
        ttcgcagatt tttcccgact ctgtaatgtt ggcggtgcag gaagggattg V S Q I F P D S V M L A V Q E G I
        acttactcac ttttccgccg gcgcccggtt ctccggagcc gcctcacctt D L L T F P P A P G S P E P P H L
251
        tcccggcagc ccgagcagcc ggagcagaga gccttgggtc cggtttctat S R Q P E Q P E Q R A L G P V S
        gccaaacctt gtaccggagg tgatcgatct tacctgccac gaggctggct M P N L V P E V I D L T C H E A G
        ttccacccag tgacgacgag gatgaagagg gtgaggagtt tgtgttagat F P P S D D E D E E G E F V L D
        tatgtggagc accccgggca cggttgcagg tcttgtcatt atcaccggag Y V E H P G H G C R S C H Y H R
        gaatacgggg gacccagata ttatgtgttc gctttgctat atgaggacct R N T G D P D I M C S L C Y M R T
        gtggcatgtt tgtctacagt cctgtgtctg aacctgagcc tgagcccgag C G M F V Y S P V S E P E P E P
        ccagaaccgg agcctgcaag acctacccgc cgtcctaaaa tggcgcctgc P E P A R P T R R P K M A P
651 tatcctgaga cgcccgacat cacctgtgtc tagagaatgc aatagtagta A I L R R P T S P V S R E C N S S
701 cggatagctg tgactccggt ccttctaaca cacctcctga gatacacccg
T D S C D S G P S N T P P E I H P
        tcgccaggct gtggaatgta tcgaggactt gcttaacgag cctgggcaac R R Q A V E C I E D L L N E P G Q
801
         ctttggactt gagctgtaaa cgccccaggc cataactcga g
p L D L S C K R P R P - XhoI
```

Figure 10. Nucleotide coding sequence of E1a cDNA (SEQ ID NO: 40). Cloning sites HindIII and XhoI, and two amino acid residue positions Y47 and C124 are underscored.

```
aagettgeeg ceaceatgag acatattate tgeeacggag gtgttattae HindII M R H I I C H G G V I
    cgaagaaatg gccgccagtc ttttggacca gctgatcgaa gaggtactgg T E E M A A S L L D Q L I E E V L
    ctgataatct tccacctcct agccattttg aaccacctac ccttcacgaa A D N L P P P S H F E P P T L H E
    ctgcatgatt tagacgtgac ggccccgaa gatcccaacg aggaggcggt L H D L D V T A P E D P N E E A
    ttcgcagatt tttcccgact ctgtaatgtt ggcggtgcag gaagggattg V S Q I F P D S V M L A V Q E G I
    acttactcac ttttccgccg gcgcccggtt ctccggagcc gcctcacctt
D L T F P P A P G S P E P P H L
    teceggeage egageagee ggageagaag geettgggte eggtttetat
    gccaaacctt gtaccggagg tgatcgatct tacctgccac gaggctggct M P N L V P E V I D L T C H E A G
    ttccacccag tgacgacgag gatgaagagg gtgaggagtt tgtgttagat F P P S D D E D E E G E E F V L D
    tatgtggagc accccgggca cggttgcagg tcttgtcatt atcaccggag Y V E H P G H G C R S C H Y H R
    gaatacgggg gacccagata ttatgtgttc gctttgctat atgaggacct R N T G D P D I M C S L C Y M R T
    gtggcatgtt tgtctacagt cctgtgtctg aacctgagcc tgagcccgag C G M F V Y S P V S E P E P E P E
    ccagaaccgg agcetgcaag acetacccgc cgtcctaaaa tggcgcctgc P P P P A R P T R R P K M A P
    tatcctgaga cgcccgacat cacctgtgtc tagagaatgc aatagtagta A I L R R P T S P V S R E C N S S
    cggatagetg tgactceggt cettetaaca caceteetga gatacaceeg
    gtggtcccgc tgtgccccat taaaccagtt gccgtgagag ttggtgggcg V V P L C P I K P V A V R V G G
    tcgccaggct gtggaatgta tcgaggactt gcttaacgag cctgggcaac R R O A V E C I E D L L N E P G Q
    ctttggactt gagctgtaaa cgccccaggc cataactcga g
P I D L S C K R P R P - XhoI
```

Figure 11. Nucleotide coding sequence of E1a\_Y47H (SEQ ID NO: 41). Cloning sites HindIII and XhoI are underscored.

```
aagcttgccg ccaccatgac catggaatct ggagcagaca accagcagag HindIII M T M E S G A D N Q Q
       tggagatgct gctgtaacag aagctgaaaa tcaacaaatg acagctcaag S G D A A V T E A E N Q Q M T A Q
       cccaaccaca gattgccaca ttagcccagg tatccatgcc agcagctcat A Q P Q I A T L A Q V S M P A A H
       gcgacatcat ctgctcccac tgtaacctta gtgcagctgc ccaatgggca A T S S A P T V T L V Q L P N G
       gacagteeaa gteeatggag ttatteagge ggeecageea teagttatte Q T V Q V H G V I Q A A Q P S V I
       agtetecaca agtecaaaca gtteagtett cetgtaagga ettaaaaaga o s p Q V Q T V Q S S C K D L K R
       cttttctccg gaactcagat ttcaactatt gcagaaagtg aggattcaca
L F S G T Q I S T I A E S E D S
       ggaatctgtg gatagtgtaa ctgattccca aaagcgaagg gaaattcttt O E S V D S V T D S Q K R R E I L
401 caaggaggcc ttcctacagg aaaattttga atgacttatc ttctgatgca S R R P S Y R K I L N D L S S D A
       ccaggggtgc caaggattga agaagaaaag tcggaagagg agacttcagc P G V P R I E E E K S E E E T S
       ccctgccatc accactgtga cagtgccaac tccgatttac cagacaagca A P A I T T V T V P T P I Y O T S
       gtgggcagta tattgccatt acccagggag gagctataca gctggctaac S G Q Y I A I T Q G G A I Q L A N
        aatggtaccg atggggtaca gggccttcag acattaacca tgaccaatgc N G T D G V Q G L Q T L T M T N
       agetgecact cageegggta ceactattet acagtatgea cagaceactg A A A T Q P G T T I L Q Y A Q T T
       atggacagca gattctagtg cccagcaacc aagttgttgt tcaagctgcc D G Q Q I L V P S N Q V V V Q A A
        tctggcgatg tacaaacata ccaaattcgt acagcacca ctagcaccat S G D V Q T Y Q I R T A P T S T
        cgccctgga gttgttatgg catcctcccc agcacttcct acgcagcctg I A P G V V M A S S P A L P T Q P
        ctgaagaagc agcccggaag agagaggttc gtctaatgaa gaacagggaa A E E A A R K R E V R L M K N R E
        gcagcaagag aatgtcgtag aaagaagaaa gaatatgtga aatgtttaga A A R E C R R K K K E Y V K C L
        gaacagagtg gcagtgcttg aaaaccaaaa caagacattg attgaggagc E N R V A V L E N Q N K T L I E E
        taaaagcact taaggacett tactgecaca aatcagatta aggatee L K A L K D L Y C H K S D - BamHI
```

Figure 12. Nucleotide coding sequence of hamster CREB-B cDNA (SEQ ID NO: 42). Cloning sites HindIII and BamHI are underscored.

```
tggagatgct gctgtaacag aagctgaaaa tcaacaaatg acagctcaag S G D A A V T E A E N Q Q M T A Q
       cccaaccaca gattgccaca ttagcccagg tatccatgcc agcagctcat A Q P Q I A T L A Q V S M P A A H
       gcgacatcat ctgctccac tgtaacctta gtgcagctgc ccaatgggca A T S S A P T V T L V Q L P N G
151
       gacagtccaa gtccatggag ttattcaggc ggcccagcca tcagttattc Q T V Q V H G V I Q A A Q P S V I
       agtctccaca agtccaaaca gttcagtctt cctgtaagga cttaaaaaaga O S P O V O T V Q S S C K D L K R
       cttttctccg gaactcagat ttcaactatt gcagaaagtg aggattcaca L F S G T Q I S T I A E S E D S
       ggaatctgtg gatagtgtaa ctgattccca aaagcgaagg gaaattcttt Q E S V D S V T D S Q K R R E I L
       caaggaggcc ttcctccagg aaaattttga atgacttatc ttctgatgca S R R P S F R K I L N D L S S D A
       ccaggggtgc caaggattga agaagaaaag tcggaagagg agacttcagc P G V P R I E E E K S E E E T S
       ccctgcatc accactgtga cagtgccaac tccgatttac cagacaagca A P A I T T V T V P T P I Y Q T S
       gtgggcagta tattgccatt acccagggag gagctataca gctggctaac S G O Y I A I T Q G G A I Q L A N
       aatggtaccg atggggtaca gggccttcag acattaacca tgaccaatgc N G T D G V O G L O T L T M T N
       agctgccact cagccgggta ccactattct acagtatgca cagaccactg A A A T O P G T T I L Q Y A Q T T
       atggacagca gattetagtg cccagcaacc aagttgttgt tcaagctgcc D G Q Q I L V P S N Q V V V Q A A
       tctggcgatg tacaaacata ccaaattcgt acagcacca ctagcaccat S G D V Q T Y Q I R T A P T S T
       cgccctgga gttgttatgg catcctcccc agcacttcct acgcagcctg I A P G V V M A S S P A L P T Q P
      ctgaagaagc agcccggaag agagaggttc gtctaatgaa gaacagggaa
A E E A A R K R E V R L M K N R E
      gcagcaagag aatgtcgtag aaagaagaaa gaatatgtga aatgtttaga A A R E C R R K K K E Y V K C L
       gaacagagtg gcagtgcttg aaaaccaaaa caagacattg attgaggagc E N R V A V L E N O N K T L I E E
       taaaagcact taaggacctt tactgccaca aatcagatta aggatcc L K A L K D L Y C H K S D - BamHI
```

Figure 13. Nucleotide coding sequence of hamster CREB-B Y134F cDNA (SEQ ID NO:

43). Cloning sites HindIII and BamHI are underscored.

1 aagcttactg ttggtaaagc cgccaccatg gaggcttggg agtgtttgga MEAWECL HindIII 51 agatttttct gctgtgcgta acttgctgga acagagctct aacagtacct EDFSAVR NLLEQSS NST cttggttttg gaggtttctg tggggctcat cccaggcaaa gttagtctgc S W F W R F L W G S S Q A K L V C agaattaagg aggattacaa gtgggaattt gaagagcttt tgaaatcctg RIKEDYKWEFEELLKS tggtgagctg tttgattctt tgaatctggg tcaccaggcg cttttccaag CGELFDSLNL G H Q A L F Q agaaggtcat caagactttg gatttttcca caccggggcg cgctgcggct 251 EKVIKTL DFS TPG RAAA gctgttgctt ttttgagttt tataaaggat aaatggagcg aagaaaccca A V A F L S F I K D K W S totgagoggg gggtacotgo tggattttot ggccatgoat otgtggagag 351 H L S G G Y L L D F L A M H L W R 401 cggttgtgag acacaagaat cgcctgctac tgttgtcttc cgtccgcccg AVVRHKNRLL LLS SVRP 451 gcgataatac cgacggagga gcagcagcag cagcaggagg aagccaggcg gcggcggcag gagcagagcc catggaaccc gagagccggc ctggaccctc 501 R R R Q E Q S P W N P R A G L D P 551 gggaatgatc taga R E - XbaI

Figure 14. Nucleotide coding sequence of E1b-19K (SEQ ID NO: 44). Cloning sites HindIII and XbaI are underscored.

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1 CCATT CCATT CALL SALES OF THE STATE OF THE
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Figure 15. Nucleotide coding sequence of hamster Bcl2 deletion mutant (SEQ ID NO: 45). Cloning sites Ncol and XbaI are underscored.

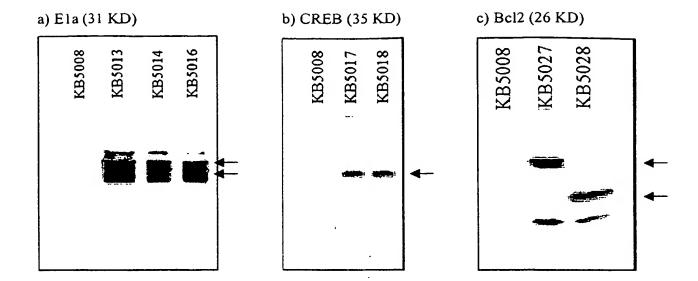


Figure 16. Western blots of E1a, CREB and Bcl2.

In a), b), and c),  $1x10^5$  CHO-K1 cells were transfected with 0.4 µg of indicated plasmid DNA. Cell lysates were collected after 48 hours and probed with appropriate antibodies. KB5008 transfected cells were used as negative controles.

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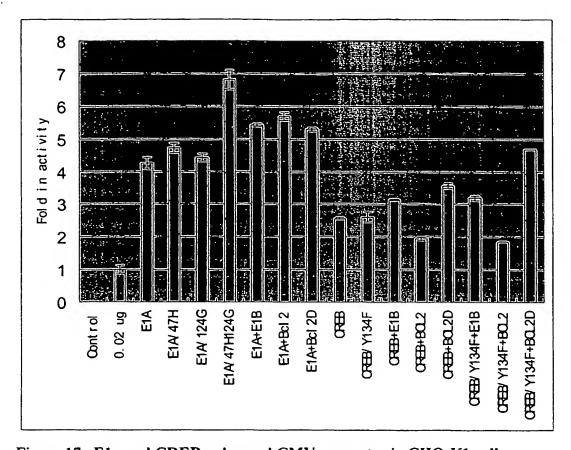


Figure 17. Ela and CREB enhanced CMV promoter in CHO-K1 cells.

 $1\times10^5$  cells were transfected with 0.02 µg of SEAP reporter construct KB5019 and 0.2 µg of E1a or CREB expressing constructs. Total DNA was added up to 0.4 µg/well by control plasmid KB5008, E1b-19K or Bcl2 expressing plasmids. SEAP activities from culture media were assay 48 hours after transfection. The control was mock transfected CHO-K1 cells.

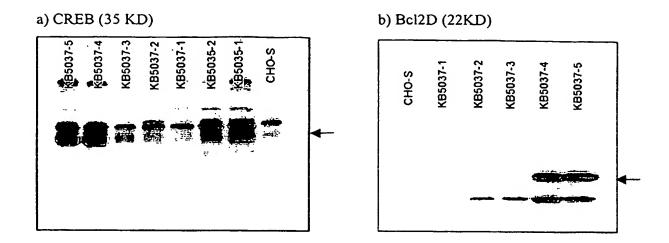


Figure 18. Western blots showing overexpression of hamster CREB-B Y134F and hamster  $\Delta$ Bcl-2 in stable transfectant CHO-S cell lines.